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L2: Entry 1 of 6

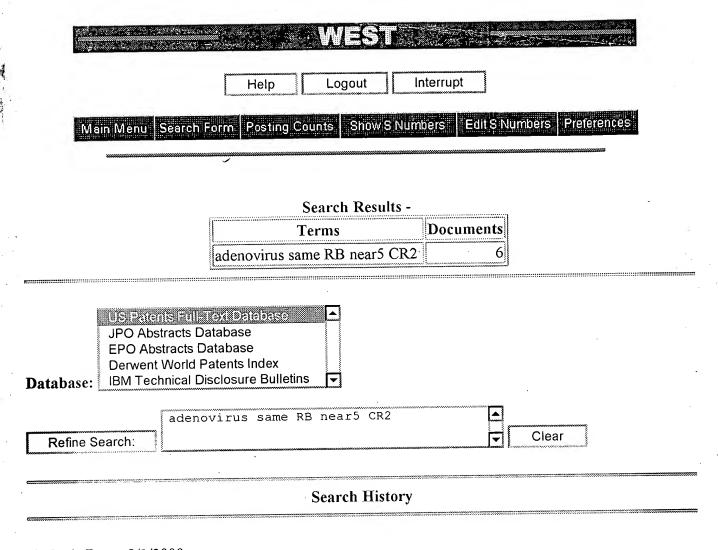
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Jun 27, 2000

DOCUMENT-IDENTIFIER: US 6080578 A TITLE: Cytopathic adenoviral E1B mutated viruses for therapy and prophylaxis of neoplasia

## BSPR:

In an alternative embodiment of the invention, a recombinant adenovirus comprising an Ela locus encoding an Ela protein (e.g., p289R or p243R) that is substantially incapable of forming a complex with RB protein in infected cells is administered to an individual or cell population comprising a neoplastic cell capable of being infected by the recombinant adenovirus. The substantial incapacity of the recombinant adenovirus to effectively sequester RB protein in infected non-neoplastic cells results in the introduced recombinant adenoviral polynucleotide(s) failing to express a replication phenotype in non-neoplastic cells. By contrast, neoplastic cells which lack a functional RB protein support expression of a replication phenotype by the introduced recombinant adenovirus which leads to ablation of the neoplastic cell by an adenoviral cytopathic effect and/or expression of a negative selection gene linked to the replication phenotype. In preferred variations of these embodiments, the recombinant adenovirus comprises an Ela locus encoding a mutant Ela protein (e.g., p289R) that lacks a CR1 and/or CR2 domain capable of binding RB (and/or the 300kD polypeptide and/or the 107kD polypeptide) but comprises a functional CR3 domain capable of transactivation of adenoviral early genes. Additional variations of these embodiments include those where the recombinant adenovirus comprises a nonfunctional Ela locus which is substantially incapable of expressing a protein that binds to and inactivates RB and may optionally also comprise a functional pl9 protein (i.e., capable of stimulating expression of adenoviral early region genes in the absence of Ela function). Recombinant adenoviruses of the invention may further comprise a mutant p19 gene which produces enhanced cytopathic effects; such a mutant known in the art is the p19 cyt mutant gene.



Today's Date: 9/1/2000

<b>DB</b> Name	<u>Query</u>	Hit Count	Set Name
USPT	adenovirus same RB near5 CR2	6	<u>L2</u>
USPT	adenovirus same E1 near5 deletion same RB near5 CR2	0	<u>L1</u>